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<110> Xenon Genetics Inc.

<120> Juvenile Hemochromatosis Gene (HFE2A), Expression Products and Uses Thereof

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<151> 2003-04-09

<150> 60/462,867

<151> 2003-04-15

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 Ala Ala Thr Ala Thr Ser Lys Leu Thr Ile Ile Phe Lys Asn Phe Gln
 210 215 220
 Glu Cys Val Asp Gln Lys Val Tyr Gln Ala Glu Met Asp Glu Leu Pro
 225 230 235 240
 Ala Ala Phe Val Asp Gly Ser Lys Asn Gly Gly Asp Lys His Gly Ala
 245 250 255
 Asn Ser Leu Lys Ile Thr Glu Lys Val Ser Gly Gln His Val Glu Ile
 260 265 270
 Gln Ala Lys Tyr Ile Gly Thr Thr Ile Val Val Arg Gln Val Gly Arg
 275 280 285
 Tyr Leu Thr Phe Ala Val Arg Met Pro Glu Glu Val Val Asn Ala Val
 290 295 300
 Glu Asp Trp Asp Ser Gln Gly Leu Tyr Leu Cys Leu Arg Gly Cys Pro
 305 310 315 320
 Leu Asn Gln Gln Ile Asp Phe Gln Ala Phe His Thr Asn Ala Glu Gly
 325 330 335
 Thr Gly Ala Arg Arg Leu Ala Ala Ala Ser Pro Ala Pro Thr Ala Pro
 340 345 350
 Glu Thr Phe Pro Tyr Glu Thr Ala Val Ala Lys Cys Lys Glu Lys Leu
 355 360 365
 Pro Val Glu Asp Leu Tyr Tyr Gln Ala Cys Val Phe Asp Leu Leu Thr
 370 375 380
 Thr Gly Asp Val Asn Phe Thr Leu Ala Ala Tyr Tyr Ala Leu Glu Asp
 385 390 395 400
 Val Lys Met Leu His Ser Asn Lys Asp Lys Leu His Leu Tyr Glu Arg
 405 410 415
 Thr Arg Asp Leu Pro Gly Arg Ala Ala Ala Gly Leu Pro Leu Ala Pro
 420 425 430
 Arg Pro Leu Leu Gly Ala Leu Val Pro Leu Leu Ala Leu Leu Pro Val
 435 440 445
 Phe Cys
 450

<210> 24
 <211> 478
 <212> PRT
 <213> Homo sapiens

<400> 24

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Met Ile Arg Lys Lys Arg Lys Arg Ser Ala Pro Pro Gly Pro Cys Arg
1           5           10           15

Ser His Gly Pro Arg Pro Ala Thr Ala Pro Ala Pro Pro Pro Ser Pro
          20           25           30

Glu Pro Thr Arg Pro Ala Trp Thr Gly Met Gly Leu Arg Ala Ala Pro
          35           40           45

Ser Ser Ala Ala Ala Ala Ala Ala Glu Val Glu Gln Arg Arg Ser Pro
          50           55           60

Gly Leu Cys Pro Pro Pro Leu Glu Leu Leu Leu Leu Leu Phe Ser
65           70           75           80

Leu Gly Leu Leu His Ala Gly Asp Cys Gln Gln Pro Ala Gln Cys Arg
          85           90           95

Ile Gln Lys Cys Thr Thr Asp Phe Val Ser Leu Thr Ser His Leu Asn
          100          105          110

Ser Ala Val Asp Gly Phe Asp Ser Glu Phe Cys Lys Ala Leu Arg Ala
          115          120          125

Tyr Ala Gly Cys Thr Gln Arg Thr Ser Lys Ala Cys Arg Gly Asn Leu
          130          135          140

Val Tyr His Ser Ala Val Leu Gly Ile Ser Asp Leu Met Ser Gln Arg
145          150          155          160

Asn Cys Ser Lys Asp Gly Pro Thr Ser Ser Thr Asn Pro Glu Val Thr
          165          170          175

His Asp Pro Cys Asn Tyr His Ser His Ala Gly Ala Arg Glu His Arg
          180          185          190

Arg Gly Asp Gln Asn Pro Pro Ser Tyr Leu Phe Cys Gly Leu Phe Gly
          195          200          205

Asp Pro His Leu Arg Thr Phe Lys Asp Asn Phe Gln Thr Cys Lys Val
          210          215          220

Glu Gly Ala Trp Pro Leu Ile Asp Asn Asn Tyr Leu Ser Val Gln Val
225          230          235          240

Thr Asn Val Pro Val Val Pro Gly Ser Ser Ala Thr Ala Thr Asn Lys
          245          250          255

Ile Thr Ile Ile Phe Lys Ala His His Glu Cys Thr Asp Gln Lys Val

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260 265 270
 Tyr Gln Ala Val Thr Asp Asp Leu Pro Ala Ala Phe Val Asp Gly Thr
 275 280 285
 Thr Ser Gly Gly Asp Ser Asp Ala Lys Ser Leu Arg Ile Val Glu Arg
 290 295 300
 Glu Ser Gly His Tyr Val Glu Met His Ala Arg Tyr Ile Gly Thr Thr
 305 310 315 320
 Val Phe Val Arg Gln Val Gly Arg Tyr Leu Thr Leu Ala Ile Arg Met
 325 330 335
 Pro Glu Asp Leu Ala Met Ser Tyr Glu Glu Ser Gln Asp Leu Gln Leu
 340 345 350
 Cys Val Asn Gly Cys Pro Leu Ser Glu Arg Ile Asp Asp Gly Gln Gly
 355 360 365
 Gln Val Ser Ala Ile Leu Gly His Ser Leu Pro Arg Thr Ser Leu Val
 370 375 380
 Gln Ala Trp Pro Gly Tyr Thr Leu Glu Thr Ala Asn Thr Gln Cys His
 385 390 395 400
 Glu Lys Met Pro Val Lys Asp Ile Tyr Phe Gln Ser Cys Val Phe Asp
 405 410 415
 Leu Leu Thr Thr Gly Asp Ala Asn Phe Thr Ala Ala Ala His Ser Ala
 420 425 430
 Leu Glu Asp Val Glu Ala Leu His Pro Arg Lys Glu Arg Trp His Ile
 435 440 445
 Phe Pro Ser Ser Gly Asn Gly Thr Pro Arg Gly Gly Ser Asp Leu Ser
 450 455 460
 Val Ser Leu Gly Leu Thr Cys Leu Ile Leu Ile Val Phe Leu
 465 470 475

<210> 25
 <211> 420
 <212> PRT
 <213> Mus musculus

<400> 25

Met Gly Gln Ser Pro Ser Pro Arg Ser Pro His Gly Ser Pro Pro Thr
 1 5 10 15
 Leu Ser Thr Leu Thr Leu Leu Leu Leu Leu Cys Gly Gln Ala His Ser
 20 25 30
 Gln Cys Lys Ile Leu Arg Cys Asn Ala Glu Tyr Val Ser Ser Thr Leu

35	40	45
Ser Leu Arg Gly Gly Gly 50	Ser Pro Asp Thr 55	Pro Arg Gly Gly Gly Arg 60
Gly Gly Leu Ala Ser 65	Gly Gly Leu Cys Arg 70	Ala Leu Arg Ser Tyr Ala 75 80
Leu Cys Thr Arg 85	Arg Thr Ala Arg Thr 90	Cys Arg Gly Asp Leu Ala Phe 95
His Ser Ala Val 100	His Gly Ile Glu Asp 105	Leu Met Ile Gln His Asn Cys 110
Ser Arg Gln Gly Pro Thr 115	Ala Pro Pro Pro 120	Ala Arg Gly Pro Ala Leu 125
Pro Gly Ala Gly Pro Ala 130	Pro Leu Thr Pro 135	Asp Pro Cys Asp Tyr Glu 140
Ala Arg Phe Ser Arg 145	Leu His Gly Arg 150	Ala Pro Gly Phe Leu His Cys 155 160
Ala Ser Phe Gly Asp 165	Pro His Val Arg 170	Ser Phe His Asn Gln Phe His 175
Thr Cys Arg Val 180	Gln Gly Ala Trp 185	Pro Leu Leu Asp Asn Asp Phe Leu 190
Phe Val Gln Ala Thr 195	Ser Ser Pro Val 200	Ser Ser Gly Ala Asn Ala Thr 205
Thr Ile Arg Lys Ile 210	Thr Ile Ile Phe Lys 215	Asn Met Gln Glu Cys Ile 220
Asp Gln Lys Val Tyr 225	Gln Ala Glu Val 230	Asp Asn Leu Pro Ala Ala Phe 235 240
Glu Asp Gly Ser 245	Ile Asn Gly Gly 250	Asp Arg Pro Gly Gly Ser Ser Leu 255
Ser Ile Gln Thr 260	Ala Asn Leu Gly 265	Ser His Val Glu Ile Arg Ala Ala 270
Tyr Ile Gly Thr Thr 275	Ile Ile Ile Arg 280	Gln Thr Ala Gly Gln Leu Ser 285
Phe Ser Ile Arg Val 290	Ala Glu Asp Val 295	Ala Arg Ala Phe Ser Ala Glu 300
Gln Asp Leu Gln Leu 305	Cys Val Gly Gly 310	Cys Pro Pro Ser Gln Arg Leu 315 320
Ser Arg Ser Glu Arg 325	Asn Arg Arg Gly 330	Ala Ile Ala Ile Asp Thr Ala 335

Arg Arg Leu Cys Lys Glu Gly Leu Pro Val Glu Asp Ala Tyr Phe Gln
 340 345 350

Ser Cys Val Phe Asp Val Ser Val Ser Gly Asp Pro Asn Phe Thr Val
 355 360 365

Ala Ala Gln Thr Ala Leu Asp Asp Ala Arg Ile Phe Leu Thr Asp Leu
 370 375 380

Glu Asn Leu His Leu Phe Pro Ser Asp Ala Gly Pro Pro Leu Ser Pro
 385 390 395 400

Ala Ile Cys Leu Val Pro Leu Leu Ser Ala Leu Phe Val Leu Trp Leu
 405 410 415

Cys Phe Ser Lys
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<210> 26

<211> 422

<212> PRT

<213> Rattus rattus

<400> 26

Met Gly Asp Arg Gly Arg Ser Pro Ser Leu Arg Ser Pro His Gly Ser
 1 5 10 15

Pro Pro Thr Leu Ser Thr Leu Thr Leu Leu Leu Leu Cys Gly Gln
 20 25 30

Ala His Ser Gln Cys Lys Ile Leu Arg Cys Asn Ala Glu Tyr Val Ser
 35 40 45

Phe Thr Leu Ser Leu Arg Gly Gly Gly Ser Pro Asp Thr Pro Arg Gly
 50 55 60

Gly Gly Arg Gly Gly Pro Ala Ser Gly Gly Leu Cys Arg Ala Leu Arg
 65 70 75 80

Ser Tyr Ala Leu Cys Thr Arg Arg Thr Ala Arg Thr Cys Arg Gly Asp
 85 90 95

Leu Ala Phe His Ser Ala Val His Gly Ile Glu Asp Leu Met Ile Gln
 100 105 110

His Asn Cys Ser Arg Gln Gly Pro Thr Ala Ser Pro Pro Ala Arg Gly
 115 120 125

Pro Ala Leu Pro Gly Ala Gly Pro Ala Pro Leu Thr Pro Asp Pro Cys
 130 135 140

Asp Tyr Glu Ala Arg Phe Ser Arg Leu His Gly Arg Thr Pro Gly Phe
 145 150 155 160

Leu His Cys Ala Ser Phe Gly Asp Pro His Val Arg Ser Phe His Asn

20

165										170					175						
His	Phe	His	Thr	Cys	Arg	Val	Gln	Gly	Ala	Trp	Pro	Leu	Leu	Asp	Asn						
			180					185					190								
Asp	Phe	Leu	Phe	Val	Gln	Ala	Thr	Ser	Ser	Pro	Val	Ala	Ser	Gly	Ala						
		195					200					205									
Asn	Ala	Thr	Thr	Ile	Arg	Lys	Ile	Thr	Ile	Ile	Phe	Lys	Asn	Met	Gln						
	210					215					220										
Glu	Cys	Ile	Asp	Gln	Lys	Val	Tyr	Gln	Ala	Glu	Val	Asp	Asn	Leu	Pro						
225					230					235					240						
Ala	Ala	Phe	Glu	Asp	Gly	Ser	Val	Asn	Gly	Gly	Asp	Arg	Pro	Gly	Gly						
				245					250					255							
Ser	Ser	Leu	Ser	Ile	Gln	Thr	Ala	Asn	Leu	Gly	Ser	His	Val	Glu	Ile						
			260					265					270								
Arg	Ala	Ala	Tyr	Ile	Gly	Thr	Thr	Ile	Ile	Val	Arg	Gln	Thr	Ala	Gly						
		275					280					285									
Gln	Leu	Ser	Phe	Ser	Ile	Arg	Val	Ala	Glu	Asp	Val	Ala	Arg	Ala	Phe						
	290					295					300										
Ser	Ala	Glu	Gln	Asp	Leu	Gln	Leu	Cys	Val	Gly	Gly	Cys	Pro	Pro	Ser						
305					310					315					320						
Gln	Arg	Leu	Ser	Arg	Ser	Glu	Arg	Asn	Arg	Arg	Gly	Ala	Ile	Ala	Ile						
				325					330					335							
Asp	Thr	Ala	Arg	Arg	Leu	Cys	Lys	Glu	Gly	Leu	Pro	Val	Glu	Asp	Ala						
			340					345					350								
Tyr	Phe	Gln	Ser	Cys	Val	Phe	Asp	Val	Ser	Val	Ser	Gly	Asp	Pro	Asn						
		355					360					365									
Phe	Thr	Val	Ala	Ala	Gln	Ser	Ala	Leu	Asp	Asp	Ala	Arg	Val	Phe	Leu						
		370				375					380										
Thr	Asp	Leu	Glu	Asn	Leu	His	Leu	Phe	Pro	Val	Asp	Ala	Gly	Pro	Pro						
385					390					395					400						
Leu	Ser	Pro	Ala	Thr	Cys	Leu	Val	Arg	Leu	Leu	Ser	Val	Leu	Phe	Val						
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Leu	Trp	Phe	Cys	Ile	Gln																
			420																		

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<212>	PRT
<213>	Fugu

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 20 25 30
 Arg Glu Ala Ala Asn Ala Glu Tyr Cys Arg Ala Leu His Ser Tyr Ser
 35 40 45
 Thr Cys Thr Lys Arg Met Ala Arg Pro Cys Arg Gly Asp Leu Ala Tyr
 50 55 60
 His Ser Ala Val Gln Gly Ile Glu Asp Leu Leu Ile Gln Tyr Arg Cys
 65 70 75 80
 Pro Leu Ala Gly Pro Thr Ala Gln Pro Arg Pro Leu Pro Pro Leu Leu
 85 90 95
 Ser Gly Asp Val Cys Leu Tyr Asp Arg Arg Leu Ala Ala Ala Glu Ala
 100 105 110
 Pro Gln Pro Asp Tyr Leu His Cys Gly Val Phe Gly Asp Pro His Ile
 115 120 125
 Arg Thr Phe Asn Asn Asp Phe His Thr Cys Ala Val Gln Gly Ala Trp
 130 135 140
 Pro Leu Ile Asp Asn Asp Phe Leu Tyr Val Gln Ala Thr Ser Ser Pro
 145 150 155 160
 Thr Arg Arg Gly Thr Gln Ala Thr Met Leu Thr Lys Ile Thr Val Ile
 165 170 175
 Val Lys Ser Trp Arg His Cys Val Asp Gln Gln Leu Tyr Gln Ala Glu
 180 185 190
 Leu Asp Asp Val Pro Met Ala Phe Ala Asp Gly Ser Val Val Ser Gly
 195 200 205
 Glu Arg Arg Gly Gln His Thr Leu Ala Ile Thr Gln Ser Pro Gly Arg
 210 215 220
 His Ala Glu Ile Arg Ala Ala His Ile Ala Thr Val Ala Ser Gly Gln
 225 230 235 240
 Ser Gly Arg Ser Leu Ser Leu Ser Val Tyr Ser Pro Arg Ser Val Val
 245 250 255
 Glu Ala Phe Gly Pro Glu Gln Asp Leu Gln Leu Cys Met Trp Gly Cys
 260 265 270
 Pro Ala Ser Gln Lys Leu Ser Thr Pro Pro Pro Thr Ser Ser Thr Phe
 275 280 285
 Ser Ala Ala Val Leu Ala His Cys Asp Ala Leu Leu Pro Val Arg Asp

290 295 300

Val Tyr His Gln Ala Cys Ile Phe Asp Leu Ile Thr Ser Gly Asp Leu
305 310 315 320

Asn Ser Ser Gly Ala Ala Ile Ser Ala Leu Gln Asp Ala Gln Lys Leu
325 330 335

Ile Ser Asp Pro Lys Arg Val His Leu Leu Ser Pro Thr Ser Ala Ala
340 345 350

Gln Arg Glu Asp His Leu Cys Leu Leu Leu Leu Leu Ser
355 360 365

<210> 28
<211> 432
<212> PRT
<213> Chicken

<400> 28
Met Gly Arg Gly Ala Gly Ser Thr Ala Leu Gly Leu Phe Gln Ile Leu
1 5 10 15

Pro Val Phe Leu Cys Ile Phe Pro Pro Val Thr Ser Pro Cys Lys Ile
20 25 30

Leu Lys Cys Asn Ser Glu Phe Trp Ala Ala Thr Ser Gly Ser His His
35 40 45

Leu Gly Ala Glu Glu Thr Pro Glu Phe Cys Thr Ala Leu Arg Ala Tyr
50 55 60

Ala His Cys Thr Arg Arg Thr Ala Arg Thr Cys Arg Gly Asp Leu Ala
65 70 75 80

Tyr His Ser Ala Val His Gly Ile Asp Asp Leu Met Val Gln His Asn
85 90 95

Cys Ser Lys Asp Gly Pro Thr Ser Gln Pro Arg Leu Arg Thr Leu Pro
100 105 110

Pro Gly Asp Ser Gln Glu Arg Ser Asp Ser Pro Glu Ile Cys His Tyr
115 120 125

Glu Lys Ser Phe His Lys His Ser Ala Ala Pro Asn Tyr Thr His Cys
130 135 140

Gly Leu Phe Gly Asp Pro His Leu Arg Thr Phe Thr Asp Thr Phe Gln
145 150 155 160

Thr Cys Lys Val Gln Gly Ala Trp Pro Leu Ile Asp Asn Asn Tyr Leu
165 170 175

Asn Val Gln Val Thr Asn Thr Pro Val Leu Pro Gly Ser Ser Ala Thr

180	185	190
Ala Thr Ser Lys Leu Thr Ile Ile Phe Lys Ser Phe Gln Glu Cys Val		
195	200	205
Glu Gln Lys Val Tyr Gln Ala Glu Met Asp Glu Leu Pro Ala Ala Phe		
210	215	220
Ala Asp Gly Ser Lys Asn Gly Gly Asp Lys His Gly Ala Asn Ser Leu		
225	230	235
Lys Ile Thr Glu Lys Val Ser Gly Gln His Ile Glu Ile Gln Ala Lys		
245	250	255
Tyr Ile Gly Thr Thr Ile Val Val Arg Gln Val Gly Arg Tyr Leu Thr		
260	265	270
Phe Ala Val Arg Met Pro Glu Glu Val Val Asn Ala Val Glu Asp Arg		
275	280	285
Asp Ser Gln Gly Leu Tyr Leu Cys Leu Arg Gly Cys Pro Leu Asn Gln		
290	295	300
Gln Ile Asp Phe Gln Thr Phe Arg Leu Ala Gln Ala Ala Glu Gly Arg		
305	310	315
Ala Arg Arg Lys Gly Pro Ser Leu Pro Ala Pro Pro Glu Ala Phe Thr		
325	330	335
Tyr Glu Ser Ala Thr Ala Lys Cys Arg Glu Lys Leu Pro Val Glu Asp		
340	345	350
Leu Tyr Phe Gln Ser Cys Val Phe Asp Leu Leu Thr Thr Gly Asp Val		
355	360	365
Asn Phe Met Leu Ala Ala Tyr Tyr Ala Phe Glu Asp Val Lys Met Leu		
370	375	380
His Ser Asn Lys Asp Lys Leu His Leu Tyr Glu Arg Thr Arg Ala Leu		
385	390	395
Ala Pro Gly Asn Ala Ala Pro Ser Glu His Pro Trp Ala Leu Pro Ala		
405	410	415
Leu Trp Val Ala Leu Leu Ser Leu Ser Gln Cys Trp Leu Gly Leu Leu		
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<211> 21

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<213> Artificial

<220>

<223> Polynucleotide replication primer

<400> 29
tccaagtcag cgactctctc g 21

<210> 30
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<220>
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<400> 30
tccaagtcag tgactctctc g 21

<210> 31
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<220>
<223> Fragment containing polymorphism

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acctgccgcg gggacctcgc c 21

<210> 32
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<220>
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<400> 32
acctgccgcg tggacctcgc c 21

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<400> 33
gcctgggaaa cctggctgga t 21

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<220>
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<400> 34
gcctgggaaa gctggctgga t 21

<210> 35
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<400> 35
tcccttctgt ctttagctca t 21

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<400> 36
tcccttctgt gttagctca t 21

<210> 37
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<220>
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<400> 37
gaggaggagg ccggggtgga 20

<210> 38
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<220>
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<400> 38
gaggaggagg aggccggggt gga 23

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<210> 39
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<400> 39
gcctccctgc cccggaccct t 21

<210> 40
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<400> 40
gcctccctgc gccggaccct t 21

<210> 41
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<400> 41
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<210> 42
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<400> 42
atggtcgtcc accgggggttc t 21

<210> 43
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<220>
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<400> 43
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<210> 44
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<400> 44
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<210> 45
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<400> 45
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<210> 46
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<212> DNA
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<220>
<223> Fragment containing polymorphism

<400> 46
gtccaaggag attggcctct a 21

<210> 47
<211> 21
<212> DNA
<213> Artificial

<220>
<223> Fragment containing polymorphism

<400> 47
cccccatggc gttgggggcc a 21

<210> 48
<211> 21
<212> DNA
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<220>
<223> Fragment containing polymorphism

<400> 48
cccccatggc tttgggggcc a 21

<210> 49
<211> 21
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<220>
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<400> 49
taagaacatg caggaatgca t 21

<210> 50
<211> 21
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<220>
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<400> 50
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<210> 51
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<400> 51
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<210> 52
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<400> 52
gccttctcag gtgaacagga c 21

<210> 53
<211> 21

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<212> DNA
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<223> Fragment containing polymorphism
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agatgctggg gttcctcttt c 21

<210> 54
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<223> Fragment containing polymorphism
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agatgctggg attcctcttt c 21

<210> 55
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<212> DNA
<213> Artificial
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<223> Forward replication primer
<400> 55
cacttgagcc caggaatttg 20

<210> 56
<211> 20
<212> DNA
<213> Artificial
<220>
<223> Reverse replication primer
<400> 56
gactcactgc agccttgacc 20

<210> 57
<211> 22
<212> DNA
<213> Artificial
<220>
<223> Forward replication primer
<400> 57
gtgtgctaca agtttgccga at 22

<210> 58
<211> 20
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<220>
<223> Reverse replication primer

<400> 58
gcttgaaact gggagttgga 20

<210> 59
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<220>
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<400> 59
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<210> 60
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<212> DNA
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<220>
<223> Reverse replication primer

<400> 60
cgccctgcca atatgttct 19

<210> 61
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<210> 62
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gtgtcacaca actggttggt